c2

60

filtered and dried under reduced pressure to provide 120 mg. of powder of 76-(4-cyano-3-methoxyisothiazol-5-yl)thio-acetamido-7x-methoxy-3-(1-methyltetrazol-5-yl)thiomethyl-cephem-4-carboxylic acid.

PS

Nuclear magnetic resonance spectra (D₆-DMSO) (p.p.m.): 3.40 (3H), 3.58 (2H), 3.92 (3H), 3.99 (3H), 4.15 (2H), 4.18 (2H), 5.12 (1H).

M

What is claimed is:

.

70. Methoxy-7 β -(4-substituted methylene-1,3-dithietan 2-yl/carboxamido-3-heterocyclic thiomethyl-\(\infty^3\)-cephem-4carboxylic acid represented by the general formula

$$\frac{R^{1}}{R^{2}} = \frac{S}{S} = CONH \frac{OCH_{3}}{OCOOH} \frac{CH_{2}-S-R^{3}}{OCOOH}$$

wherein R represents a carboxyl group or the functional derivative residue thereof; R2 represents a hydrogen atom, a lower alkyl group, a lower alkoxy group, R4S(0), group (wherein R4 represents a lower alkyl group and n represents 0, 1 or 2), a lower Alkanoyl group, an aryl group, an aroyl group, a carboxyl group or the functional derivative residue thereof, a lower alkenyl group, a sulfamoyl group, or a heterocyclic residue; and R3 represents a lower alkyl-substituted tetrazolyl group or a lower alkyl-substituted thiadiazofyl group

d the phermaceutically acceptable salts thereof.

216 GO

- 62558 g 7β - {4-[(Carbamoyl)(carboxy)methylene]-1,3-dithietan,-2-yl}carboxamido-7x-methoxy-3-(1-methyltetrazol -5-yl)thiomethyl- \triangle^3 -cephem-4-carboxylic acid as claimed in claim 1.
 - $7\beta-[4-(1-Carboxyethylidene)-1,3-dithietane-2-yl]$ carboxamido-7~methoxy-3-(1-methyltetrazol -5-yl)thiomethyl- \triangle^3 -cephem-4-carboxylic acid as claimed in claim 1.
 - $7\beta \{4 \{(Carboxy) (methoxy) methylene\} 1, 3 dithietan_{A} = 1, 3 dithietan_{A} =$ 2-yl}carboxamido-7<-methoxy-3-(1-methyltetrazol -5-yl)thio- \downarrow methyl- \triangle 3-cephem-4-carboxylic acid as claimed in claim 1.

- 5. 7β -{4-[(Carboxy)(methylthio)methylene]-1,3-dithietan,-2-yl}carboxamido-7 α -methoxy-3-(l-methyltetrazol -5-yl)thio-methyl- Δ 3-cephem-4-carboxylic acid as claimed in claim l.
- 6. $7\beta \{4 [(Carboxy)(ethylthio)methylene] 1, 3 dithietan_ 2 yl\} carboxamido 7\lambda methylene 2 yl) thio-methyl-\(\Delta^3 cephem 4 carboxylic acid as claimed in claim 1.$
 - 7. 7β-[4-(Carboxymethylene)-1,3-dithietan-2-yl)carboxamido-7 -methoxy-3-(1-methyltetrazol -5-yl)thiomethyl-Δ³-cephem-4-) carboxylic acid as claimed in claim 1.
 - 8. A process for the preparation of 7χ -methoxy- 7β =

 (4-substituted methylene-1,3-dithietan-2-yl)carboxamido-3heterocyclic thiomethyl- Δ^3 -cephem-4-carboxylic acid
 represented by the general formula

$$\frac{R^{1}}{R^{2}} = \frac{S}{S} - CONH - \frac{OCH_{3}}{O} - CH_{2} - S - R^{2}$$

wherein R¹, R² and R³ have the same significance as/claim 1, which comprises reacting the 4-substituted methylene-1,3-dithietan-2-carboxylic acid represented by the general formula

wherein R^1 and R^2 have the same significance as above, or the functional derivative thereof, with the 7β -amino- 7χ -methoxy-3-heterocyclic thiomethyl- Δ^3 -cephem-4-carboxylic acid represented by the general formula

107-107

K

wherein R3 has the same significance as above.

9. A process for the preparation of %-methoxy- 7β (4-substituted methylene-1,3-dithietan-2-yl)carboxamido3-heterocyclic thiomethyl- Δ^3 -cep_hem-4-carboxylic acid
represented by the general formula

wherein R^1 , R^2 and R^3 have the same significance as in claim 1, which comprises reacting the 3-acetoxymethyl- (or 3-carbamoyl-oxymethyl-) 7%-methoxy-7β-(4-substituted methylene-1,3-dithietan-2-yl)carboxamido- Δ^3 -cephem-4-carboxylic acid represented by the general formula

wherein R¹ and R have the same significance as above and R⁶ represents an acetyl group or a carbamoyl group,
with the heterocyclic thiol represented by the general formula

$$HS - R^3$$

wherein R3/ has the same significance as above.

Jyr. J

wherein R² and R³ have the same significance as in claim 1 and R⁸ represents a hydrogen atom or a substituted or unsubstituted alkyl group,

under a basic condition which comprises treating/the //x-methoxy-3-heterocyclic thiomethylcephalosporin derivative represented by the

general formula

wherein R^2 , R^3 and R^8 have the same significance as above.

 7β -(3-Hydroxy-4-substituted isothiazol-5-yl)thio-acetamido-7 χ -methoxy-3-heterocyclic thiomethyl- Δ -cephem-4-carboxylic acid represented by the general formula

wherein R² and R³ have the same significance as in claim 1. and the pharmaceutically acceptable non-toxic salts thereof.

9 12. 78-(4-Carboxy-3-hydroxyisothiazol-5-yl)thioacetamido-74-methoxy-3-(1-methyltetrazol-5-yl)thiomethyl- Δ^3 -cephem-4-carboxylic acid as claim in claim 12.

End-

110-